File Copy 09/105117

(FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, SCISEARCH' ENTERED AT 18:57:06 ON 19 OCT 2001 2893 S MICROBIAL (W) PRODUCTION L1L2 190 S L1 AND (AMINO (W) ACIDS) L3 28 S L2 AND CORYNEBACTERIUM L413 L3 AND LYSINE O L3 AND ((EXPORT) (W) (GENE OR CARRIER)) L6 0 S L3 AND EXPORT (W) GENE L7 166 S EXPORT (W) GENE 0 S L3 AND L7 L8 L9 0 S L3 (P) L7 0 S L2 AND L7 L10 0 S L2 AND EXPORT (W) GENE L1161 S L7 AND MICROB? 1 S L12 AND CORYNEBACTERIUM 26 DUP REM L3 (2 DUPLICATES REMOVED) 13 DUP REM L4 (0 DUPLICATES REMOVED)

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STN Search Shategy

File Copy 09/105117 EAST Search Strategy

E ro rs	0	0	0	0				
Error Definition								
Comment								
Time Stamp	2001/10/1 9 18:38	2001/10/1 9 18:38	2001/10/1 9 18:39	2001/10/1 9 18:40				
DBs	USPA T; US-P GPUB; JPO; JPO; DERW	USPA T; US-P GPUB; JPO; JPO; DERW	USPA T; US-P GPUB; JPO; JPO; DERW	USPA T; US-P GPUB; JPO; JPO;				
Search Text	microbial adj production adj of adj amino adj acid	microbial adj production	L7 and (amino adj acid)	L13 and Corynebacterium				
Hits	0	702	314	47				
T #	L1	L.7	L13	L1 0				
Type	BRS	BRS	BRS	BRS				
	1	7	м	4				

10/19/2001, EAST Version: 1.02.0008

Type L # Hits Se	L # Hits	Hits	Se	Search Text	DBs	Time Stamp	Comment s	Error Definition	Er or rs
BRS L25 0 (manufacturing a L-1ysine)	L19 and (manufacturing L-lysine)	L19 and (manufacturing L-lysine)		adj	USPA T; US-P GPUB ; JPO; JPO; DERW	2001/10/1 9 18:42		-	0
L19 and ((export) adj (gene carrier))	П		L19 and ((export)adj(ger carrier))	le or		2001/10/1 9 18:47			0
BRS L31 11 L19 and lysine	11 L19 and	1 L19 and	and		USPA T; US-P GPUB ; JPO; DERW ENT	2001/10/1 9 18:48	`		0

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FILE 'SCISEARCH' ENTERED AT 18:57:06 ON 19 OCT 2001
COPYRIGHT (C) 2001 Institute for Scientific Information (ISI) (R)
=> s microbial (w) production
          2893 MICROBIAL (W) PRODUCTION
L1
=> s L1 and (amino (w) acids)
           190 L1 AND (AMINO (W) ACIDS)
=> s L2 and Corynebacterium
            28 L2 AND CORYNEBACTERIUM
=> L3 and lysine
            13 L3 AND LYSINE
L4
=> L3 and ((export) (w) (gene or carrier))
             0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))
=> s L3 and export (w) gene
             0 L3 AND EXPORT (W) GENE
=> s export (w) gene
1.7
           166 EXPORT (W) GENE
=> s L3 and L7
             0 L3 AND L7
L8
=> s L3 (p) L7
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L13 (P) L37'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L14 (P) L38'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L15 (P) L39'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L16 (P) L40'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L17 (P) L41'
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0 L3 (P) L7

0.15

```
\stackrel{'}{=} s L2 and L7
             0 L2 AND L7
=> s L2 and export (w) gene
             0 L2 AND EXPORT (W) GENE
=> s L7 and microb?
L12
            61 L7 AND MICROB?
=> s L12 and Corynebacterium
L13
             1 L12 AND CORYNEBACTERIUM
=> dup rem L3
PROCESSING COMPLETED FOR L3
             26 DUP REM L3 (2 DUPLICATES REMOVED)
=> dup rem L4
PROCESSING COMPLETED FOR L4
T.15
             13 DUP REM L4 (0 DUPLICATES REMOVED)
=> dis L13 ibib kwic
L13 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER:
                         1997:475788 CAPLUS
DOCUMENT NUMBER:
                         127:172154
                         L-Lysine export from Corynebacterium
TITLE:
                         glutamicum. Physiological and molecular-biological
                         characterization of the carrier-mediated export of a
                         primary metabolite
AUTHOR(S):
                         Vrljic, Maria-Marina
CORPORATE SOURCE:
                         Inst. Biotechnologie, Forschungszentrum Julich
                         G.m.b.H., Juelich, D-52425, Germany
SOURCE:
                         Ber. Forschungszent. Juelich (1997), Juel-3349, 1-115
                         pp.
                         CODEN: FJBEE5; ISSN: 0366-0885
DOCUMENT TYPE:
                         Report
LANGUAGE:
                         German
     L-Lysine export from Corynebacterium glutamicum. Physiological
     and molecular-biological characterization of the carrier-mediated export
     of a primary metabolite
AΒ
     The gene for the Lys-excretion carrier was isolated from C. glutamicum
and
     the Lys export was analyzed physiol. A system was established which
     induces the Lys excretion in dependence of Met. The mutant NA8, defect
in
     Lys export, was isolated. The L-Lys export (LysE) gene encodes a
     polypeptide of 236 amino acids with the potential to span the membrane 6
     times and a mol. wt. of 2,5425 Da. With overexpressed LysE, L-Lys was
     exported at a rate of 3.76 nmol/min/mg dry wt. which lead to a 10-fold
     increased Lys excretion rate. The LysG (governing L-Lys export)
     gene is localized immediately adjacent to LysE, but is
     transcripted divergently. The deduced polypeptide (290 amino acids) has
     helix-turn-helix motive at the aminoterminus. At the sequence level,
LysG
     shows .ltoreq.35% identity to prokaryotic, autoregulatory transcriptional
```

```
regulators. LysG acts in trans and leads to a decrease of the Lys
     excretion by C. glutamicum. For the Lys-export defect mutant C.
     glutamicum NA8, the transition G1594.fwdarw.A1594 was shown which results
     in a stop-codon in the LysE gene. The resulting LysE polypeptide in C.
     glutamicum NA8 is shortened for 43 amino acids. The growth of a LysEG
     deletion mutant was abolished on a minimal medium in the presence of
     Lys-contg. dipeptides. The quantification of the intracellular L-Lys
     concns. revealed an accumulation of Lys .ltoreq.1,100 mM. The results
     suggest that the physiol. function of the Lys export carrier of C.
     glutamicum is to avoid extremely high intracellular Lys concns.
ST
     lysine excretion carrier Corynebacterium gene sequence; protein
     sequence Corynebacterium lysine excretion carrier
IT
     Amino acid transport (biological)
        (carrier-mediated, export; lysine export from Corynebacterium
        glutamicum, carrier-supported export of a primary metabolite)
IT
     Helix-turn-helix
        (gene lysG protein; lysine export from Corynebacterium
        glutamicum, carrier-supported export of a primary metabolite)
     Proteins (specific proteins and subclasses)
TT
     RL: BAC (Biological activity or effector, except adverse); BOC
(Biological
     occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)
        (gene lysG, (governing lysine export); lysine export from
        Corynebacterium glutamicum, carrier-supported export of a
        primary metabolite)
IT
     Genes (microbial)
     RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological
     study); OCCU (Occurrence)
        (lysE; lysine export from Corynebacterium glutamicum,
        carrier-supported export of a primary metabolite)
IT
     Genes (microbial)
     RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological
     study); OCCU (Occurrence)
        (lysG (governing lysine export); lysine export from
       Corynebacterium glutamicum, carrier-supported export of a
       primary metabolite)
IT
    Corynebacterium glutamicum
     DNA sequences
     Protein sequences
        (lysine export from Corynebacterium glutamicum,
       carrier-supported export of a primary metabolite)
ΙT
    Amino acid transporters
     RL: BAC (Biological activity or effector, except adverse); BOC
(Biological
    occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)
        (lysine-transporting, gene lysE; lysine export from
       Corynebacterium glutamicum, carrier-supported export of a
       primary metabolite)
IT
     63-68-3, L-Methionine, biological studies
     RL: BAC (Biological activity or effector, except adverse); BIOL
     (Biological study)
        (induces lysine excretion; lysine export from Corynebacterium
       glutamicum, carrier-supported export of a primary metabolite)
     184922-77-8, GenBank X96471-derived protein GI 1729755
    RL: BAC (Biological activity or effector, except adverse); BOC
(Biological
    occurrence); PRP (Properties); BIOL (Biological study); OCCU (Occurrence)
        (lysine export from Corynebacterium glutamicum,
```

```
carrier-supported export of a primary metabolite)
     184922-76-7, GenBank X96471-derived protein GI 1729754
IT
                                                               184922-78-9,
     GenBank X96471-derived protein GI 1729756
     RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological
     study); OCCU (Occurrence)
        (lysine export from Corynebacterium glutamicum,
        carrier-supported export of a primary metabolite)
IT
     56-87-1, L-Lysine, biological studies
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (lysine export from Corynebacterium glutamicum,
        carrier-supported export of a primary metabolite)
     184343-19-9, GenBank X96471
IT
     RL: BOC (Biological occurrence); PRP (Properties); BIOL (Biological
     study); OCCU (Occurrence)
        (nucleotide sequence; lysine export from Corynebacterium
        glutamicum, carrier-supported export of a primary metabolite)
=> dis L14 1-26 ibib kwic
L14 ANSWER 1 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS
ACCESSION NUMBER:
                    2001:314459 BIOSIS
DOCUMENT NUMBER:
                    PREV200100314459
TITLE:
                    Effect of gluconic acid as a secondary carbon source on
                    non-growing L-lysine producers cells of
                    Corynebacterium glutamicum. Purification and
                    properties of 6-phosphogluconate dehydrogenase.
AUTHOR(S):
                    Bianchi, Daniella; Bertrand, Olivier; Haupt, Karsten;
                    Coello, Nereida (1)
CORPORATE SOURCE:
                    (1) Instituto de Biologia Experimental, Universidad
Central
                    deVenezuela, Caracas, 1041-A: ncoello@uole.com.ve
Venezuela
SOURCE:
                    Enzyme and Microbial Technology, (June 7, 2001) Vol. 28,
                    No. 9-10, pp. 754-759. print.
                    ISSN: 0141-0229.
DOCUMENT TYPE:
                    Article
LANGUAGE:
                    English
SUMMARY LANGUAGE:
                    English
     Effect of gluconic acid as a secondary carbon source on non-growing
     L-lysine producers cells of Corynebacterium glutamicum.
     Purification and properties of 6-phosphogluconate dehydrogenase.
AB
     We studied the production of L-lysine in Corynebacterium
     glutamicum ATCC 21543 non growing cells obtained by nutrient limitation.
     Statistical analysis revealed significant differences in the L-lysine
     titers of.
IT
        Engineering; Methods and Techniques; Nutrition
IT
     Chemicals & Biochemicals
        6-phosphogluconate dehydrogenase: amino acid sequence, analysis,
        molecular properties, pH, purification; L-lysine: microbial
       production, yield; amino acids: analysis;
        carbon sources; gluconic acid: secondary carbon source
ORGN .
       Microorganisms; Irregular Nonsporing Gram-Positive Rods: Actinomycetes
        and Related Organisms, Eubacteria, Bacteria, Microorganisms;
       Microorganisms
ORGN Organism Name
```

Bacillus subtilis (Endospore-forming Gram-Positives); Corynebacterium glutamicum (Irregular Nonsporing Gram-Positive Rods): non-growing cells; Escherichia coli (Enterobacteriaceae); bacteria (Bacteria); microorganisms (Microorganisms) ORGN Organism Superterms Bacteria; Eubacteria; Microorganisms L14 ANSWER 2 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS 2001:421002 BIOSIS ACCESSION NUMBER: DOCUMENT NUMBER: PREV200100421002 TITLE: L-glutamate fermentation and metabolic engineering: Studies on the L-glutamate production mechanism in Coryneform bacteria. AUTHOR(S): Nakamatsu, Tsuyoshi Nippon Nogeikagaku Kaishi, (Jun., 2001) Vol. 75, No. 6, SOURCE: pp. 683-686. print. ISSN: 0002-1407. DOCUMENT TYPE: General Review LANGUAGE: Japanese SUMMARY LANGUAGE: English Major Concepts Biochemistry and Molecular Biophysics; Bioprocess Engineering; Metabolism TΤ Chemicals & Biochemicals amino acids: large-scale microbial production; glutamate: large-scale microbial production; oxoglutarate dehydrogenase ORGN Super Taxa Irregular Nonsporing Gram-Positive Rods: Actinomycetes and Related Organisms, Eubacteria, Bacteria, Microorganisms ORGN Organism Name Corynebacterium spp. (Irregular Nonsporing Gram-Positive Rods) ORGN Organism Superterms Bacteria; Eubacteria; Microorganisms L14 ANSWER 3 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS ACCESSION NUMBER: 2001:174925 BIOSIS DOCUMENT NUMBER: PREV200100174925 TITLE: MALDI-TOF MS for quantification of substrates and products in cultivations of Corynebacterium glutamicum. AUTHOR(S): Wittmann, Christoph (1); Heinzle, Elmar CORPORATE SOURCE: (1) Biochemical Engineering Institute, Saarland University, 66041, Saarbruecken: c.wittmann@rz.uni-sb.de Germany SOURCE: Biotechnology and Bioengineering, (March 20, 2001) Vol. 72, No. 6, pp. 642-647. print. ISSN: 0006-3592. DOCUMENT TYPE: Article LANGUAGE: English

cultivations of Corynebacterium glutamicum.

English

MALDI-TOF MS for quantification of substrates and products in

SUMMARY LANGUAGE:

Major Concepts

ΙT

Biochemistry and Molecular Biophysics; Bioprocess Engineering; Methods and Techniques

IT Chemicals & Biochemicals

amino acids: microbial production

, quantitative analysis; products: quantitative analysis; substrates: quantitative analysis

L14 ANSWER 4 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 20

2000:761605 CAPLUS

DOCUMENT NUMBER:

134:99608

TITLE:

Development and use of miniaturized parallel experiment technology for bioprocess development

AUTHOR(S):

Altenbach-Rehm, Jutta

CORPORATE SOURCE:

Institut fur Biotechnologie, Julich, JUL-3782,

Germany SOURCE:

Ber. Forschungszent. Juelich (2000), Juel-3782, a-f,

i-iv, 1-233

CODEN: FJBEE5; ISSN: 0366-0885

DOCUMENT TYPE: LANGUAGE:

Report German

AB The fed-batch technique is nowadays the std. operation mode for high performance microbial prodn. processes. Shake flasks are widely used a simple bioreactors in batch process development. Because of uncontrolled changes in pH and reduced oxygen transfer rates parallel operated shake flasks can usually not be applied for microbial fed-batch process development. Due to the need to operate controlled stirred tank reactors the development of specific fermn. strategies is expensive and time consuming. To address these issues a miniature fed-batch technique was developed in cooperation with DASGIP mbH, Julich, and INFORS AG, Basel. Controlled batch and fed-batch fermns. can be performed in 16 parallel small scale bioreactors. The new technique allows the feeding of up to 4 different substrates and parallel pH control

based on user-defined profiles. The feeding assembly is sterilized chem. using dimethylcarbonate. To overcome the limitations of shake flasks, small scale bubble columns were developed. Gas distribution is performed with a new type of sterile sparger. Transferring fed-batch fermns. from small scale bubble columns to a stirred tank reactor a scale up factor of 20 was achieved. Escherichia coli K12 was chosen to test the new parallel

bioreactor technique. Compared to shake flask fermns, the cell concn. was

50% higher due to efficient oxygen transfer. In fed-batch fermns. with pH-controlled substrate feeding up to 35 g/l DCW were achieved. For the 1st time, a parallel optimization of feeding profiles in parallel small scale fed-batch expts. with successful scale-up to a lab. bioreactor was performed. Escherichia coli BL 21 (DE3) pLysS produces the recombinant enzyme GDP-.alpha.-D-mannose-pyrophosphorylase after induction with isopropyl-.beta.-D-thiogalactoside (IPTG). Single induction with 0,5 mM IPTG resulted in a low specific enzyme activity of 1,6 U/g DCW (dry cell wt.). For the optimization of enzyme expression a genetic algorithm was used. A final enzyme activity of 111 U/g DCW was achieved for optimal substrate and inducer feeding profiles. To demonstrate the advantages of this new parallel bioreactor technique different strains of industrial relevance were investigated. Corynebacterium glutamicum, Staphylococcus carnosus and Ashbya gossypii.

ST bioreactor bubble column fed batch fermn; GDP mannose pyrophosphorylase bubble column fed batch; Staphylococcus bubble column fed batch fermn;

```
isoleucine bubble column fed batch Corynebacterium; riboflavin
     bubble column fed batch Ashbya
IT
     Amino acids, biological studies
     RL: BPR (Biological process); BIOL (Biological study); PROC (Process)
        (amino acid consumption in riboflavin prodn. by Ashybya gossypii in
        parallel bubble columns with fed-batch technique)
IT
     Corynebacterium glutamicum
        (L-isoleucine prodn. by Corynebacterium glutamicum in
        parallel bubble columns with fed-batch technique)
IT
     73-32-5P, L-Isoleucine, biological studies
     RL: BMF (Bioindustrial manufacture); BPR (Biological process); BIOL
     (Biological study); PREP (Preparation); PROC (Process)
        (L-isoleucine prodn. by Corynebacterium glutamicum in
        parallel bubble columns with fed-batch technique, amino acid
        consumption in riboflavin prodn. by Ashybya gossypii)
TΤ
     61-90-5, L-Leucine, biological studies
     RL: BPR (Biological process); MFM (Metabolic formation); BIOL (Biological
     study); FORM (Formation, nonpreparative); PROC (Process)
        (L-isoleucine prodn. by Corynebacterium glutamicum in
        parallel bubble columns with fed-batch technique, amino acid
        consumption in riboflavin prodn. by Ashybya gossypii)
L14 ANSWER 5 OF 26 CAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER:
                        1999:244776 CAPLUS
DOCUMENT NUMBER:
                        130:266420
TITLE:
                        Method for microbial production of
                         amino acids of the aspartate and/or
                         glutamate family and agents which can be used in said
                        method
INVENTOR(S):
                        Eikmanns, Bernd; Peters-Wendisch, Petra; Sahm,
Hermann
PATENT ASSIGNEE(S):
                         Forschungszentrum Julich G.m.b.H., Germany
SOURCE:
                         PCT Int. Appl., 40 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         German
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
    PATENT NO.
                    KIND DATE
                                         APPLICATION NO. DATE
                    ____
                                          ______
                      A2
    WO 9918228
                           19990415
                                         WO 1998-EP6210 19980930
    WO 9918228
                      A3
                           19990520
        W: AU, BR, CA, CN, HU, ID, JP, KR, MX, RU, SK, US
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE
    DE 19831609
                      A1
                            19990415
                                          DE 1998-19831609 19980714
    AU 9911482
                      A1
                            19990427
                                          AU 1999-11482
                                                           19980930
    EP 1015621
                           20000705
                      Α2
                                          EP 1998-954301
                                                           19980930
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, FI
                                          BR 1998-13021
    BR 9813021
                      Α
                           20000815
                                                           19980930
PRIORITY APPLN. INFO.:
                                       DE 1997-19743894 A 19971004
                                       DE 1998-19831609 A 19980714
                                       WO 1998-EP6210
                                                       W 19980930
TΤ
    Method for microbial production of amino
```

acids of the aspartate and/or glutamate family and agents which

can be used in said method

AB The invention relates to a method for microbial prodn. of amino acids of the aspartate and/or glutamate family in which the pyruvate carboxylase activity is increased by genetically changing the enzyme and/or the pyruvate carboxylase gene

expression of a microorganism which produces the corresponding amino acid.

agents which can be used in the inventive method.
amino acid fermn Corynebacterium pyruvate carboxylase genetic

ST amino acid fermn Corynebacterium pyruvate carboxylase genetic engineering

IT Corynebacterium glutamicum

Fermentation

(microbial prodn. of amino acids

of the aspartate and/or glutamate family and agents which can be used in said method)

In addn., the invention relates to a pyruvate carboxylase gene and addnl.

IT Amino acids, preparation

RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)

(microbial prodn. of amino acids

of the aspartate and/or glutamate family and agents which can be used in said method)

IT Genetic engineering

(microbial prodn. of amino acids

of the aspartate and/or glutamate family and modification of **Corynebacterium** pyc gene in said method)

IT Genes (microbial)

RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (pyc; microbial prodn. of amino

acids of the aspartate and/or glutamate family and modification
of Corynebacterium pyc gene in said method)

IT 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 72-19-5P, L-Threonine, preparation 74-79-3P, L-Arginine, preparation 672-15-1P, L-Homoserine

RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation)

(microbial prodn. of amino acids

of the aspartate and/or glutamate family and agents which can be used in said method)

IT 9014-19-1, Pyruvate carboxylase 204116-18-7 208541-81-5

RL: BSU (Biological study, unclassified); BIOL (Biological study)

(microbial prodn. of amino acids

of the aspartate and/or glutamate family and agents which can be used in said method)

L14 ANSWER 6 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 1999:175414 BIOSIS DOCUMENT NUMBER: PREV199900175414

TITLE: Cloning of the transketolase gene and the effect of its

dosage on aromatic amino acid production in

Corynebacterium glutamicum.

AUTHOR(S): Ikeda, M. (1); Okamoto, K.; Katsumata, R.

CORPORATE SOURCE: (1) Technical Research Laboratories, Kyowa Hakko Kogyo

Co.,

Ltd., Hofu, Yamaguchi, 747-8522 Japan

SOURCE: Applied Microbiology and Biotechnology, (Feb., 1999) Vol.

51, No. 2, pp. 201-206.

ISSN: 0175-7598.

DOCUMENT TYPE:

Article

LANGUAGE: English

 ${\tt TI}$ Cloning of the transketolase gene and the effect of its dosage on aromatic

amino acid production in Corynebacterium glutamicum.

AB. . . enzyme of the non-oxidative pentose phosphate pathway. The effect of

its overexpression on aromatic amino acid production was investigated in <code>Corynebacterium</code> glutamicum, a typical amino-acid-producing organism. For this purpose, the transketolase gene of the organism was cloned on the basis of. . . the presence of the gene in high copy numbers enabled tyrosine, phenylalanine and tryptophan producers to accumulate 5%-20% more aromatic <code>amino acids</code>. These results indicate that overexpressed transketolase activity operates to redirect the glycolytic intermediates toward the nonoxidative pentose phosphate pathway in. . .

IT Major Concepts

Bioprocess Engineering; Enzymology (Biochemistry and Molecular Biophysics); Molecular Genetics (Biochemistry and Molecular Biophysics)

IT Chemicals & Biochemicals

aromatic amino acids: microbial

production; transketolase [EC 2.2.1.1]; Corynebacterium
transketolase gene (Irregular Nonsporing Gram-Positive Rods)

L14 ANSWER 7 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1998:277651 CAPLUS

DOCUMENT NUMBER:

128:307587

TITLE:

Microbial production of substances

from aromatic metabolism

INVENTOR(S):

Sprenger, Georg; Siewe, Ruth; Sahm, Hermann; Karutz,

Martin; Sonke, Theodorus

PATENT ASSIGNEE(S):

Forschungszentrum Juelich G.m.b.H., Germany; Holland

Sweetener Co. V.o.F.

SOURCE:

Ger. Offen., 14 pp. CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND				ND	DATE APPL					CATI	и ис	ο.	DATE				
DE	1964	 4566		 A	 1	1998	0430			E 19	 96-1	9644	 566	1996	 1026		
WO	9818	936		A	1	1998	0507		W	o 19	97-N	L582		1997	1017		
	W:	AL,	ΑU,	BA,	BB,	BG,	BR,	CA,	CN,	CU,	CZ,	EE,	GE,	HU,	ID,	IL,	IS,
		JP,	KP,	KR,	LC,	LK,	LR,	LT,	LV,	MG,	MK,	MN,	MX,	NO,	NZ,	PL,	RO,
		SG,	SI,	SK,	SL,	TR,	TT,	UA,	US,	UZ,	VN,	YU,	AM,	ΑZ,	BY,	KG,	KZ,
		MD,	RU,	TJ,	TM												
	RW:	GH,	ΚE,	LS,	MW,	SD,	SZ,	UG,	ZW,	AT,	BE,	CH,	DE,	DK,	ES,	FI,	FR,
		GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,
		GN,	ML,	MR,	ΝE,	SN,	TD,	TG									
AU	AU 9747277 A1			1998	0522												
EP	9344	18		A.	1	1999	0811		E.	P 19	97-9	0974	8	1997	1017		
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	IT,	LI,	NL,	SE,	PT,	FI		
CN	1241	214		Α		2000	0112		Cl	N 19	97-1	8090	8	1997	1017		
JP	2001	5064	86	\mathbf{T}^{2}	2	2001	0522		J.	P 19	98-5	2031	8	1997	1017		
PRIORIT	Y APP	LN.	INFO	. :					DE 1:	996-	1964	4566	Α	1996	1026		
								1	WO 1	997-1	NL58:	2	W	1997:	1017		

```
ΤI
     Microbial production of substances from aromatic
     metabolism
IT
     Transport proteins
     RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU
     (Biological use, unclassified); BIOL (Biological study); OCCU
     (Occurrence); PREP (Preparation); USES (Uses)
        (gene glf glucose facilitator protein, of Zymomonas mobilis;
        microbial prodn. of substances from arom. metab.)
IΤ
     Genes (microbial)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (glf, for glucose facilitator protein of Zymomonas mobilis;
        microbial prodn. of substances from arom. metab.)
TΤ
     Genes (microbial)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (glk, for glucokinase of Zymomonas mobilis; microbial
        prodn. of substances from arom. metab.)
ΙT
     Pentose phosphate pathway
        (intermediates of, in amino acid manuf.; microbial
        prodn. of substances from arom. metab.)
ΙT
     Bacillus (bacterium genus)
     Brevibacterium
       Corynebacterium
     Escherichia
     Escherichia coli
     Fermentation
     Microorganism
     Molecular cloning
     Serratia
        (microbial prodn. of substances from arom. metab.)
TТ
     Transport proteins
     RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU
     (Biological use, unclassified); BIOL (Biological study); OCCU
     (Occurrence); PREP (Preparation); USES (Uses)
        (microbial prodn. of substances from arom. metab.)
ΙT
     Amino acids, preparation
     RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP
     (Preparation)
        (microbial prodn. of substances from arom. metab.)
TΤ
     Plasmids
        (pZ4557tal; microbial prodn. of substances from
        arom. metab.)
IT
     Plasmids
        (pZ4557tkt; microbial prodn. of substances from
        arom. metab.)
IT
     Plasmids
        (pZ4557tkttal; microbial prodn. of substances from
        arom. metab.)
ΙT
     Genes (microbial)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (talB; microbial prodn. of substances from arom.
       metab.)
TΤ
     Genes (microbial)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (tktA; microbial prodn. of substances from arom.
```

```
metab.)
ΙT
     9001-36-9P, Glucokinase
     RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU
     (Biological use, unclassified); BIOL (Biological study); OCCU
     (Occurrence); PREP (Preparation); USES (Uses)
        (gene glk, of Zymomonas mobilis; microbial prodn.
        of substances from arom. metab.)
IT
     585-18-2, Erythrose-4-phosphate
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (in amino acid manuf.; microbial prodn. of
        substances from arom. metab.)
     9014-46-4P, Transaldolase
IT
                                9014-48-6P, Transketolase
     RL: BOC (Biological occurrence); BPN (Biosynthetic preparation); BUU
     (Biological use, unclassified); BIOL (Biological study); OCCU
     (Occurrence); PREP (Preparation); USES (Uses)
        (microbial prodn. of substances from arom. metab.)
     63-91-2P, L-Phenylalanine, preparation
IT
     RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP
     (Preparation)
        (microbial prodn. of substances from arom. metab.)
L14 ANSWER 8 OF 26 CAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER:
                        1995:674124 CAPLUS
DOCUMENT NUMBER:
                         123:54314
TITLE:
                        Enhancement of reduced NADP production for enhanced
                        microbial production of biochemicals
INVENTOR(S):
                        Kojima, Hiroyuki; Totsuka, Kazuhiko
PATENT ASSIGNEE(S):
                        Ajinomoto Co., Inc., Japan
                        PCT Int. Appl., 32 pp.
SOURCE:
                        CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         Japanese
FAMILY ACC. NUM. COUNT:
                        1
PATENT INFORMATION:
     PATENT NO.
                    KIND DATE
                                         APPLICATION NO. DATE
                     ---- -----
     _____
     WO 9511985
                          19950504
                      A1
                                         WO 1994-JP1791
                                                           19941026
         W: AU, BR, CA, CN, CZ, HU, JP, KR, PL, RU, SK, US, VN
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
     CA 2175042
                            19950504
                      AA
                                          CA 1994-2175042 19941026
    AU 9480026
                                           AU 1994-80026
                      A1
                            19950522
                                                            19941026
    AU 687458
                      В2
                            19980226
     EP 733712
                      Α1
                            19960925
                                          EP 1994-931158
                                                          19941026
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT,
SE
     BR 9407907
                      Α
                            19961126
                                           BR 1994-7907
                                                            19941026
     HU 74840
                      A2
                            19970228
                                           HU 1996-1085
                                                          19941026
     ZA 9503350
                      Α.
                            19961025
                                           ZA 1995-3350
                                                            19950425
                            19981103
     US 5830716
                      Α
                                           US 1996-619521
                                                           19960429
     CN 1139956
                      A
                            19970108
                                           CN 1994-194707
                                                            19961026
PRIORITY APPLN. INFO.:
                                        JP 1993-270828
                                                            19931028
                                        WO 1994-JP1791
                                                            19941026
     Enhancement of reduced NADP production for enhanced microbial
TΤ
    production of biochemicals
```

AB The productivity of such substances as L-amino acids, antibiotics, vitamins, growth factors and physiol. active substances in

the fermn. using a microorganism is improved by improving the productivity of reduced NADP in the cells of the microorganisms. Construction of pMW::THY contg. the Escherichia coli transhydrogenase gene, and introduction of the plasmid into the L-threonine-producing Escherichia coli B-3996 were shown. The recombinant E. coli B-3996 produced L-threonine .apprx.10% higher than did the parental strain. ΙT Corynebacterium glutamicum Escherichia coli Fermentation (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) TΤ Amino acids, preparation RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) IT Plasmid and Episome (pHSG::THY; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) IT Plasmid and Episome (pMW::THY; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) ΙT Plasmid and Episome (pSU::THY; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) IT 9014-18-0, Nicotinamide nucleotide transhydrogenase RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study) (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) ΤТ 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 61-90-5P, L-Leucine, preparation 63-91-2P, L-Phenylalanine, preparation 72-18-4P, L-Valine, preparation 72-19-5P, L-Threonine, preparation 73-32-5P, L-Isoleucine, preparation RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) IT 53-59-8P, NADP RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (reduced; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) L14 ANSWER 9 OF 26 SCISEARCH COPYRIGHT 2001 ISI (R) ACCESSION NUMBER: 95:184306 SCISEARCH THE GENUINE ARTICLE: OK574 TITLE: METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM CORYNEBACTERIUM-GLUTAMICUM AUTHOR: SAHM H (Reprint); EGGELING L; EIKMANNS B; KRAMER R CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL, D-52425 JULICH, GERMANY (Reprint) GERMANY COUNTRY OF AUTHOR: SOURCE: FEMS MICROBIOLOGY REVIEWS, (FEB 1995) Vol. 16, No. 2-3, pp. 243-252. ISSN: 0168-6445.

Article; Journal

DOCUMENT TYPE:

FILE SEGMENT: LIFE LANGUAGE: ENGLISH

REFERENCE COUNT:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM CORYNEBACTERIUM -GLUTAMICUM

The Gram-positive bacterium Corynebacterium glutamicum is AΒ used for the industrial production of amino acids, e.g. of L-glutamate and L-lysine. In the last 10 years, genetic engineering and amplification of relevant structural genes have become.

STAuthor Keywords: CORYNEBACTERIUM GLUTAMICUM; AMINO ACID PRODUCTION; METABOLIC DESIGN; L-LYSINE; L-THREONINE; L-ISOLEUCINE

STP KeyWords Plus (R): L-THREONINE; L-LYSINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; MICROBIAL PRODUCTION; RESISTANT MUTANTS; SPLIT PATHWAY; BIOSYNTHESIS; ISOLEUCINE; GENES

L14 ANSWER 10 OF 26 SCISEARCH COPYRIGHT 2001 ISI (R)

ACCESSION NUMBER:

96:7187 SCISEARCH

THE GENUINE ARTICLE: TJ545

METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM TITLE:

CORYNEBACTERIUM-GLUTAMICUM

AUTHOR: SAHM H (Reprint)

KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL, CORPORATE SOURCE:

D-52425 JULICH, GERMANY (Reprint)

COUNTRY OF AUTHOR: GERMANY

FOLIA MICROBIOLOGICA, (1995) Vol. 40, No. 1, pp. 23-30. SOURCE:

> ISSN: 0015-5632. Article; Journal

FILE SEGMENT: LIFE; AGRI ENGLISH LANGUAGE:

REFERENCE COUNT: 26

DOCUMENT TYPE:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

TΙ METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM CORYNEBACTERIUM-GLUTAMICUM

The Gram-positive bacterium Corynebacterium glutamicum is AB

used for the industrial production of amino acids, e.g. of L-glutamate and L-lysine. By cloning and expressing the various genes of the L-lysine pathway in C. glutamicum we. .

KeyWords Plus (R): L-THREONINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; MICROBIAL PRODUCTION; LYSINE BIOSYNTHESIS; RESISTANT MUTANTS; SPLIT PATHWAY; GENES; AMPLIFICATION; FERMENTATION

L14 ANSWER 11 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1985:436156 CAPLUS

DOCUMENT NUMBER:

103:36156

TITLE:

AUTHOR(S):

Optimization of amino acid production by automatic

self-tuning digital control of redox potential Radjai, Mohammad K.; Hatch, Randolph T.; Cadman,

Theodore W.

CORPORATE SOURCE:

Dep. Chem. Nucl. Eng., Univ. Maryland, College Park,

MD, 20742, USA

SOURCE: Biotechnol. Bioeng. Symp. (1984), 14 (Symp.

Biotechnol.

Fuels Chem., 6th), 657-79 CODEN: BIBSBR; ISSN: 0572-6565 DOCUMENT TYPE: Journal LANGUAGE: English

AB The microbial prodn. of homoserine [672-15-1], lysine

[56-87-1], and valine [72-18-4] by an auxotrophic mutant of **Corynebacterium** glutamicum was investigated in a 16-L batch fermentor. Closed-loop digital control of redox potential was implemented

using proportional-integral (PI) control of agitation rates. Due to the nonlinearity of the system, the PI controller parameters had to be changed

during the course of the fermns. An automatic, self-tuning algorithm was developed for stable control of redox potential. This permitted exptl. optimization of total and selective amino acid prodn. Total amino acid yields of 35% from glucose were achieved compared to 23% reported in the literature for the same fermn.

IT Corynebacterium glutamicum

(amino acid manuf. with, optimization and redox potential control in)

IT Amino acids, preparation

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)

(manuf. of, by fermn.)

L14 ANSWER 12 OF 26 CAPLUS COPYRIGHT 2001 ACS DUPLICATE 1

ACCESSION NUMBER:

1979:20814 CAPLUS

DOCUMENT NUMBER:

90:20814

TITLE:

Microbial production of essential

amino acids With

Corynebacterium glutamicum mutants

AUTHOR(S): CORPORATE SOURCE: Nakayama, Kiyoshi; Araki, Kazumi; Kase, Hiroshi Tokyo Res. Lab., Kyowa Hakko Kogyo Co., Ltd.,

Machida,

Japan

SOURCE:

Adv. Exp. Med. Biol. (1978), 105 (Nutr. Improv. Food

Feed Proteins), 649-61

CODEN: AEMBAP; ISSN: 0065-2598

DOCUMENT TYPE:

LANGUAGE:

Journal English

TI Microbial production of essential amino

acids with Corynebacterium glutamicum mutants

Amino acids produced by microbial processes are generally L-forms. The stereospecificity of the amino acids produced by fermn. makes the process advantageous compared with synthetic processes. Microorganisms employed in microbial processes for amino acid prodn. are divided into 4 classes: wild-type, auxotrophic mutant, regulatory mutant, and auxotrophic regulatory mutant. Using such mutants of Corynebacterium glutamicum, all the essential amino acids but L-methionine are now being produced by direct fermn. from cheap C sources such as carbohydrate materials or acetic acid.

- ST amino acid manuf Corynebacterium
- IT Corynebacterium glutamicum

(amino acid manuf. by)

IT Fermentation

(amino acids, by Corynebacterium glutamicum)

IT Amino acids, preparation

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)

(manuf. of, from carbohydrates by Corynebacterium glutamicum)

L14 ANSWER 13 OF 26 CAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 1976:521806 CAPLUS DOCUMENT NUMBER: 85:121806 TITLE: Microbial production of amino acid INVENTOR(S): Tsuchida, Takayasu; Yoshihara, Yasuhiko; Kubota, Koji; Hirose, Yoshio PATENT ASSIGNEE(S): Ajinomoto Co., Inc., Japan SOURCE: Japan. Kokai, 5 pp. CODEN: JKXXAF Patent DOCUMENT TYPE: LANGUAGE: Japanese FAMILY ACC. NUM. COUNT: 1 PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE _________ -----JP 51061690 A2 19760528 JP 1974-134879 19741122 Microbial production of amino acid amino acid manuf Brevibacterium; Corynebacterium amino acid manuf Brevibacterium TΤ Corynebacterium (amino acid manuf. by) TΨ Fermentation (amino acids, by Corynebacterium or Brevibacterium) 56-45-1P, preparation ΙT 73-22-3P, preparation RL: PREP (Preparation) (by fermn., with Corynebacterium) L14 ANSWER 14 OF 26 CAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 1975:529947 CAPLUS DOCUMENT NUMBER: 83:129947 TITLE: Microbial production of amino acids. VI. Formation of Lamino acids from DL-.alpha.-hydroxycarboxylic acids Matsushima, Hirochika; Murata, Keijiro; Mase, Yasuo AUTHOR(S): CORPORATE SOURCE: Ferment. Res. Lab., Sankyo Co., Ltd., Tanashi, Japan SOURCE: Hakko Kogaku Zasshi (1975), 53(7), 443-9 CODEN: HKZAA2 DOCUMENT TYPE: Journal LANGUAGE: Japanese Microbial production of amino acids . VI. Formation of L-amino acids from DL-.alpha.-hydroxycarboxylic acids AΒ Formation of L-amino acids from DL-.alpha.hydroxycarboxylic acids was studied. L-.alpha.-aminobutyric acid [1492-24-6] was formed in a medium contg. DL-.alpha.-hydroxybutyric acid [600-15-7] by various bacteria belonging to Aerobacter, Bacillus, Corynebacterium, Escherichia, Flavobacterium, Micrococcus, Proteus, Pseudomonas, Sarcina, Staphylococcus, and other genera. A. cloacae IAM 1221 was cultured in a medium contg. DL-.alpha.-bromobutyric acid [2385-70-8] (hydrolyzed to hydroxybutyric acid). L-.alpha.-aminobutyric acid was isolated from the culture broth and

identified by thin-layer chromatog., elementary anal., and by its specific

rotation and IR spectrum. Formation of valine [72-18-4], leucine [61-90-5], or phenylalanine [63-91-2] from DL-.alpha.-hydroxycarboxylic acids by Brevibacterium roseum ATCC 13825 was studied. Yields (mole) from

the cultures were 84.22, 95.7, and 47.7%, resp. An amino-group donor (glutamic acid) was needed besides the bacterial cells and DL-.alpha.-hydroxycarboxylic acid for the enzymic formation of amino acids.

L14 ANSWER 15 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1975:137747 CAPLUS

DOCUMENT NUMBER:

82:137747

TITLE:

Microbial production of

amino acids

INVENTOR(S):

Kubota, Koji; Yoshihara, Yasuhiko; Okada, Hiroshi

PATENT ASSIGNEE(S):

Ajinomoto Co., Inc.

SOURCE:

Japan. Kokai, 5 pp. CODEN: JKXXAF

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
JP 49109585	A2	19741018	JP 1973-24049	19730228
JP 51038796	B4	19761023		

TI Microbial production of amino acids

Amino acids were produced by a microbe cultured in a propionic acid medium. Thus, Brevibacterium flavum ATCC 14,067, Micrococcus glutamicus ATCC 13,032, Corynebacterium acetoacidophilum ATCC 13,870, Microbacterium ammoniaphilum ATCC 15,354, and B. flavum FERM-P 1684 were cultured with shaking at 31.degree. for 48 hr in a medium (pH 7.5) contg. propionic acid 2, (NH4)2SO4 1, KH2PO4 0.1, MgSO4.cntdot.7H2O 0.04, NaCl 0.1, and soybean protein hydrolysate (total N

= 7%) 0.2% plus biotin 2 and thiamine.cntdot.HCl 200 .mu.g/l. Prodn. of L-glutamic acid by each organism was 4.3, 4.2, 3.9, 4.0, and 2.5 mg/ml, resp. B. flavum FERM-P 1684 also produced N-acetylglutamine at 0.4 mg/ml.

IT Corynebacterium acetoacidophilum

(glutamic acid manuf. by, from propionic acid)

L14 ANSWER 16 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1973:56392 CAPLUS

DOCUMENT NUMBER:

78:56392

TITLE:

Microbial production of

amino acids from aromatic compounds.

I. Screening of aromatic compound-assimilating

bacteria

AUTHOR(S):

Yamamoto, Masao; Nishida, Hiroshi; Inui, Taiji;

Ozaki,

Asaichiro

CORPORATE SOURCE:

Cent. Res. Lab., Sanraku-Ocean Co., Ltd., Fujisawa,

Japan

SOURCE:

Hakko Kogaku Zasshi (1972), 50(12), 868-75

CODEN: HKZAA2

DOCUMENT TYPE: LANGUAGE:

Journal English

Microbial production of amino acids

from aromatic compounds. I. Screening of aromatic compound-assimilating bacteria

AΒ In an attempt to produce amino acids from aromatic compds. by fermn., bacterial stock cultures in this lab. were examd. for their assimilability of benzoate and salicylate; 96 strains from 97 glutamate-producing cultures assimilated benzoic acid. Then, 10 type-strains of the glutamate-producing strains were tested for their assimilability of 40 aromatic compds. 16 of the compds. were assimilated. These were benzoic acid, m-hydroxybenzoic acid, p-hydroxybenzoic acid, 2,4-dihydroxybenzoic acid, 2,5-dihydroxybenzoic acid, 3,4-dihydroxybenzoic acid, 3,5-dihydroxybenzoic acid, phenylacetic acid, phenylpyruvic acid, .beta.-phenylpropionic acid, cinnamic acid, benzal dehyde, benzyl alc., phenol, catechol, and resorcinol. A sizable amt. of L-glutamic

was produced from the assimilated compds. by these glutamate-producing bacteria, benzoate, esp., serving as the best substrate.

IT Brevibacterium

Brevibacterium lactofermentum

Corynebacterium acetoglutamicum

Microbacterium ammoniaphilum

Micrococcus glutamicus

(glutamic acid formation by, from arom. compds.)

L14 ANSWER 17 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1971:84222 CAPLUS

DOCUMENT NUMBER:

74:84222

TITLE:

Utilization of hydrocarbons by microorganisms. XXI.

Biochemical studies of microbial production of .alpha.-ketoglutarate,

L-glutamate, and DL-alanine from hydrocarbons

AUTHOR(S):

Imada, Yukio; Yamada, Koichi

CORPORATE SOURCE: SOURCE:

Fac. Agric., Univ. Tokyo, Tokyo, Japan Agr. Biol. Chem. (1971), 35(1), 18-26

CODEN: ABCHA6

DOCUMENT TYPE: Journal LANGUAGE: English

Utilization of hydrocarbons by microorganisms. XXI. Biochemical studies of microbial production of .alpha.-ketoglutarate, L-glutamate, and DL-alanine from hydrocarbons

AΒ Strain S10B1 of Corynebacterium hydrocarboclastus produced .alpha.9ketoglutaric acid (I), LGlutamate, and DLAlanine from nAlkanes in a thiam (II)Limited medium supplemented with Fe2+. The replacement of hydrocarbon substrate by sugars such as glucose not only decreased the yields, but also reversed the order of the yields among the 3 products. This phenomenon was explained by a metabolic pathway in relation to the role of II. Slow O uptake in the presence of pyruvate and I by IIDeficient cells supported the presumption that II limitation resulted

in

deficiency of a cofactor in the enzymic oxidn. of pyruvate and I. Activities of terminal enzymes in the synthesis of LGlutamate and DLLanine

were detd. and discussed. Three intermediates were detected in the culture broth.

ST Corynebacterium ketoglutarate prodn; ketoglutarate prodn Corynebacterium; glutamate prodn Corynebacterium;

alanine prodn Corynebacterium; thiamine Corynebacterium

; hydrocarbons ultilization bacteria; bacteria hydrocarbons utilization

ITCorynebacterium

(hydrocarboclastus, amino acids formation by, from

hydrocarbons)

ΙT 59-43-8, biological studies RL: BIOL (Biological study)

(amino acids formation from paraffins by

Corynebacterium hydrocarboclastus in response to)

L14 ANSWER 18 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1970:475660 CAPLUS

DOCUMENT NUMBER:

73:75660

TITLE:

Microbial production of L-glutamic

PATENT ASSIGNEE(S):

Asahi Chemical Industry Co., Ltd.

SOURCE:

Fr. Demande, 11 pp.

CODEN: FRXXBL

DOCUMENT TYPE:

Patent

LANGUAGE:

French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ----------

FR 2009795 19700206

PRIORITY APPLN. INFO.:

JΡ 19680531

Microbial production of L-glutamic acid

L-Glutamic acid (I) is prepd. by aerobic cultivation of Corynebacterium or Brevibacterium in liq. media contg. C1-3 alcs. as C source and penicillin. Thus, B. vitalumen var propanolophilum ATCC 21391 was grown in a medium contg. PrOH 50, corn steep liquor 4, KH2PO4 2,

MgSO4.7H2O 0.5, Fe2+ 0.01, Mn2+ 0.01, urea 4 g/l., with the addn. of 100 .mu.q biotin and penicillin G (K salt) 10 units/1., at 32.degree. and pH 6.5-8.0 with shaking for 96 hr to give 23.1 g I/l. (46.2% based on PrOH). PrOH and penicillin were added in portions during the fermentation. Without penicillin addn., the yield was 6.4% I.

Brevibacterium glutamate prodn; glutamate prodn Brevibacterium;

amino acids Corynebacterium;

Corynebacterium amino acids; penicillin

bacteria glutamate

IT Corynebacterium

(melassecola and petrophylum, glutamic acid manuf. by)

L14 ANSWER 19 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1970:508239 CAPLUS

DOCUMENT NUMBER:

73:108239

TITLE:

Microbial production of

L-threonine

INVENTOR(S): PATENT ASSIGNEE(S):

Nakayama, Kiyoshi; Kase, Hiroshi Kyowa Fermentation Industry Co. Ltd.

SOURCE: Ger. Offen., 22 pp.

CODEN: GWXXBX

DOCUMENT TYPE: LANGUAGE:

Patent German

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

DE 1817666 A 19700827 DE 1968-1817666 19681224

TI Microbial production of L-threonine

AB Various microorganisms, e.g. Aerobacter [Enterobacter] aerogenes, Serratia

marcescens, or Arthrobacter paraffineus, cultured for producing L-threonine required 2 or 3 of the **amino acids** isoleucine, methionine, lysine, or diaminopimelic acid. The microorganisms were cultured aerobically in an aq. medium contg. the optimal (or less) amts. of the required **amino acids**. Thus, E. aerogenes NM-IS-5 (ATCC 21,215) was cultured 96 hr at 30.degree. in medium contg. glucose 5, (NH4)2SO4 1.4, KH2PO4 0.05, K2HPO4 0.05, MgSO4.7H2O 0.025, FeSO4.7H2O 0.001, MnSO4.4H2O 0.001, and CaCO3 2% and isoleucine 50, methionine 100, and diaminopimelic acid 200 mg/l. to give 7.8 g L-threonine/l.

ST microbial prodn threonine; threonine microbial prodn; Aerobacter threonine fermn; amino acid prodn fermn

IT Corynebacterium

(glutamicum, threonine manuf. by)

L14 ANSWER 20 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER: 1970:401150 CAPLUS

DOCUMENT NUMBER: 73:1150

TITLE: Microbial production of

L-threonine. II. Production by

.alpha.-amino-.beta.-

hydroxyvaleric acid resistant mutants of glutamate

producing bacteria

AUTHOR(S): Shiio, Isamu; Nakamori, Shigeru

CORPORATE SOURCE: Cent. Res. Lab., Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE: Agr. Biol. Chem. (1970), 34(3), 448-56

CODEN: ABCHA6

DOCUMENT TYPE:

LANGUAGE:

English

TI Microbial production of L-threonine. II. Production

Journal

by .alpha.-amino-.beta.-hydroxyvaleric acid resistant mutants of glutamate

producing bacteria

AB A mutant strain of Brevibacterium flavum was able to grow in a medium contg. 5 mg DL-threo-.alpha.-amino-.beta.-hydroxyvaleric acid (AHV)/ml; 1 mg AHV/ml inhibited the growth of the parental strain by >90%. Further treatment of the AHV-resistant strain with the mutagen,

N-methyl-N'-nitro-N-nitrosoguanidine, produced a bacterial strain that was

able to grown on 8 mg AHV/ml; this mutant produced 13.5 g L-threonine/l., an amt. 30% more than that produced by the parental strain. A similarly derived mutant of **Corynebacterium** acetoacidophilum resistant to AHV produced 6.1 g threonine/l. Other **amino acids** biosynthesized by the bacteria were discussed in relation to the regulation of threonine synthesis.

ST threonine prodn bacterial; corynebacterium threonine prodn; Brevibacterium threonine prodn; mutations bacteria threonine; bacteria mutations threonine; aminohydroxyvalerate bacteria

IT Corvnebacterium

(acetoacidophilum, tryptophan formation from aminohydroxyvaleric acid

L14 ANSWER 21 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER: 1970:123860 BIOSIS

DOCUMENT NUMBER:

BA51:33860

TITLE:

MICROBIAL PRODUCTION OF AMINO

-ACIDS FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY CORYNEBACTERIUM-HYDROCARBOCLASTUS-R-

AUTHOR(S):

SHIIO I; UCHIO R

SOURCE:

AMINO ACID NUCLEIC ACID, (1969) (19), 88-96.

CODEN: HATAA4. ISSN: 0517-6174.

FILE SEGMENT:

BA; OLD

LANGUAGE:

Unavailable

MICROBIAL PRODUCTION OF AMINO-ACIDS

FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY

CORYNEBACTERIUM-HYDROCARBOCLASTUS-R-7.

L14 ANSWER 22 OF 26 BIOSIS COPYRIGHT 2001 BIOSIS DUPLICATE 2

ACCESSION NUMBER: 1970:106213 BIOSIS

DOCUMENT NUMBER:

BA51:16213

TITLE:

MICROBIAL PRODUCTION OF AMINO

-ACIDS FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY CORYNEBACTERIUM-HYDROCARBOCLASTUS

R-7.

AUTHOR(S):

SHIIO I; UCHIO R

SOURCE:

J GEN APPL MICROBIOL, (1969) 15 (1), 65-84.

CODEN: JGAMA9. ISSN: 0022-1260.

FILE SEGMENT:

BA; OLD

LANGUAGE:

Unavailable

MICROBIAL PRODUCTION OF AMINO-ACIDS

FROM HYDRO CARBONS PART 4 L GLUTAMIC-ACID PRODUCTION BY

CORYNEBACTERIUM-HYDROCARBOCLASTUS R-7.

L14 ANSWER 23 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1967:514494 CAPLUS

DOCUMENT NUMBER:

67:114494

TITLE:

Microbial production of

amino acids from hydrocarbons. III.

L-Ornithine production by an arginine auxotrophic

mutant of Corynebacterium hydrocarboclastus Ishu, Ryosuke; Ishii, Ryosuke; Shiio, Isamu Ajinomoto Co., Inc., Kawasaki, Japan

AUTHOR(S):

CORPORATE SOURCE: SOURCE:

J. Gen. Appl. Microbiol. (1967), 13(3), 3303-12

CODEN: JGAMA9

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Microbial production of amino acids

from hydrocarbons. III. L-Ornithine production by an arginine auxotrophic mutant of Corynebacterium hydrocarboclastus

AΒ cf. CA 67: 89718u. The arginine auxotrophic mutant strain RN-362 of C. hydrocarboclastus R-7 was used to study L-ornithine productiion from hydrocarbons, in a fermentation medium contq. various n-alkanes. L-Ornithine production required L-arginine at the optimum level of

q./l. of medium; an excess inhibited the biosynthesis of L-ornithine. (NH4)2HPO4 was the best source of N and, at 2% in a neutral to slightly acidic pH, gave the highest level of L-ornithine production and cell

growth; NH4OAc, KNO3, and (NH4)2CO3 proved less suitable because of a drop in pH along with the accumulation of a large amt. of .alpha.-ketoglutaric acid, pyruvic acid, and proline in the growth medium. Of 17 C sources, n-tetradecane best supported cell growth and L-ornithine production and the other C13-C17 n-alkanes did so moderately, while kerosene and light

oil produced good cell growth but only a small amt. of L-ornithine.

Addn.

of 3 g. yeast ext. and 0.5 g. L-arginine-HCl to 1 l. of medium enhanced L-ornithine production. A similar effect was achieved by replacing the yeast ext. with various amino acids at 0.01% in the medium. L-Methionine was most effective for the production of L-ornithine, while L-lysine, L-cysteine, L-cystine, L-histidine, and L-phenylalanine were less so, in decreasing order. Amino acids enhance L-ornithine production by stimulating hydrocarbon oxidn. and cell growth.

ST HYDROCARBONS USE BACTERIA; BACTERIA HYDROCARBONS USE; ALKANES USE BACTERIA; **AMINO ACIDS** PRODN HYDROCARBONS; ORNITHINE PRODN HYDROCARBONS; PARAFFINS UTILIZATION BACTERIA

IT Corynebacterium

(hydrocarboclastus, ornithine formation from hydrocarbons by)

IT Hydrocarbons, biological studies

RL: BIOL (Biological study)

(ornithine formation from, by Corynebacterium

hydrocarboclastus)

IT 70-26-8

RL: FORM (Formation, nonpreparative)

(formation of, from hydrocarbons by Corynebacterium

hydrocarboclastus)

L14 ANSWER 24 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1967:489718 CAPLUS

DOCUMENT NUMBER:

CORPORATE SOURCE:

67:89718

TITLE:

Microbial production of

amino acids from hydrocarbons. II.

Isolationf good hycarbon utilizers and amino acid

production by their auxotrophs

AUTHOR(S):

Ishii, Ryosuke; Otsuka, Shinichiro; Shiio, Isamu Central Res. Labs., Ajinomoto Co., Inc., Kawasaki,

Japan

SOURCE:

J. Gen. Appl. Microbiol. (1967), 13(2), 217-25

CODEN: JGAMA9

DOCUMENT TYPE:

Journal English

LANGUAGE: English
TI Microbial production of amino acids

from hydrocarbons. II. Isolationf good hycarbon utilizers and amino acid

production by their auxotrophs

AB cf. CA 59: 14313h. Nine microorganisms, which showed good growth on long-chain aliphatic hydrocarbons, were isolated by an enrichment culture method, followed by a single colony isolation technique. They included 5 strains of Alcaligenes marshallii, 2 strains of Corynebacterium hydrocarboclastus, and 2 strains of yeast. Various auxotrophic mutants were derived from these microorganisms. The mutants accumulated the following amino acids from aliphatic hydrocarbons; L-ornithine, L-valine, L-glutamic acid, L-leucine, L-tyrosine, L-alanine, L-proline, L-aspartic acid, and L-lysine.

ST BACTERIA AMINO ACID PRODN; AMINO ACID PRODN BACTERIA; HYDROCARBONS

AMINO ACIDS; ALIPHATICS BACTERIA METAB

IT Corynebacterium

(hydrocarboclastus, amino acid fermentation of hydrocarbons by)

ΤТ Amino acids, preparation

> RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP (Preparation)

(manuf. of, by fermentation of hydrocarbons)

L14 ANSWER 25 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1966:22870 CAPLUS

DOCUMENT NUMBER:

64:22870

ORIGINAL REFERENCE NO.: 64:4230g-h,4231a

TITLE:

Microbial production of

nucleotides

INVENTOR(S):

Masuo, Eitaro; Okabayashi, Tadashi

PATENT ASSIGNEE(S):

Shionogi & Co., Ltd.

SOURCE:

10 pp.

DOCUMENT TYPE:

Patent

LANGUAGE:

Unavailable

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

APPLICATION NO. DATE PATENT NO. KIND DATE -----JP 40010957 19650601 JP 19591214

Microbial production of nucleotides TТ

AB Some bacteria strains of high nucleotide-forming activity were detected based on the results of the test developed by the authors, and compns. of media for promoting accumulation of nucleotides were also investigated. To evaluate the nucleotide-forming activity of bacteria, cells of nonexacting purine (I) auxotrophic mutant B 96 of Escherichia coli were mixed into the synthetic medium contg. no I for testing strains. The activity of nucleotide accumulation of the strains increased as the

of the mutant increased. By this procedure, the following strains were found to be suitable for nucleotide production: Bacillus subtilis IFO 3061, B. firmus IFO 3330, B. circulans IFO 3342, B. megaterium IFO 3003, Alcaligenes viscosus AN-14, A. metalcaligenes 1021, Serratia marcescens 1008, S. plymuthica IFO 3055, Bacterium ketoglutaricum 1041, and new species of Brevibacterium and Corynebacterium. For promoting nucleotide production with these strains, amino acids, esp. L-glutamic acid (II), are necessary in the medium. Proteins or peptides contg. II are also effective for the strains having sufficient protease. Sufficient content of PO43- at pH 5.0-7.5 is also necessary

for

the medium. By cultivation under these conditions, AMP, CDP, UMP, and UDP

are obtained.

L14 ANSWER 26 OF 26 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1963:476777 CAPLUS

DOCUMENT NUMBER:

59:76777

ORIGINAL REFERENCE NO.: 59:14313h,14314a

TITLE:

Microbial production of

amino acids from hydrocarbons. I.

Preliminary screening of glutamic acid-producing

bacteria

AUTHOR(S):

Shiio, Isamu; Otsuka, Shinichiro; Ishii, Ryosuke;

Katsuya, Nobu; Iizuka, Hiroshi

CORPORATE SOURCE: Ajinomoto Co., Inc., Kawasaki, Japan

SOURCE:

J. Gen. Appl. Microbiol. (Tokyo) (1963), 9, 23-30

DOCUMENT TYPE: LANGUAGE:

Journal Unavailable

Microbial production of amino acids

from hydrocarbons. I. Preliminary screening of glutamic acid-producing

bacteria

Various bacteria utilized kerosene, light oil, heavy oil, and liquid AΒ paraffin as the only C source for growth and formation of L-glutamic acid (I). The highest level of I (281 .gamma./ml.) was obtained from kerosene by a strain of Corynebacterium hydrocarboclastus.

=> dis L15 1-13 ibib kwic

L15 ANSWER 1 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS

ACCESSION NUMBER:

2001:314459 BIOSIS

DOCUMENT NUMBER:

PREV200100314459

TITLE:

Effect of gluconic acid as a secondary carbon source on

non-growing L-lysine producers cells of Corynebacterium glutamicum. Purification and

properties of 6-phosphogluconate dehydrogenase.

AUTHOR(S):

Bianchi, Daniella; Bertrand, Olivier; Haupt, Karsten;

Coello, Nereida (1)

CORPORATE SOURCE:

(1) Instituto de Biologia Experimental, Universidad

Central

deVenezuela, Caracas, 1041-A: ncoello@uole.com.ve

Venezuela

SOURCE:

Enzyme and Microbial Technology, (June 7, 2001) Vol. 28,

No. 9-10, pp. 754-759. print.

ISSN: 0141-0229.

DOCUMENT TYPE: Article LANGUAGE: English SUMMARY LANGUAGE: English

Effect of gluconic acid as a secondary carbon source on non-growing Llysine producers cells of Corynebacterium glutamicum.

Purification and properties of 6-phosphogluconate dehydrogenase.

AB We studied the production of L-lysine in Corynebacterium glutamicum ATCC 21543 non growing cells obtained by nutrient limitation. Statistical analysis revealed significant differences in the Llysine titers of glucose, gluconic acid or glucose-gluconic acid cultures. Higher L-lysine titer obtained in batch cultures with mixed carbon sources or gluconic acid alone were found to be associated . . dehydrogenase activity (6PGDH, E.C.1.1.1.44). This enzyme is a pivotal enzyme within the hexose monophosphate pathway, and thus of importance for L-lysine production. 6PGDH was purified and characterized. The purified enzyme migrates as a single band on sodium dodecyl sulfate-polyacrylamide gel electrophoresis.

IT Bioprocess Engineering; Methods and Techniques; Nutrition

Chemicals & Biochemicals TT6-phosphogluconate dehydrogenase: amino acid sequence, analysis, molecular properties, pH, purification; L-lysine:

microbial production, yield; amino

acids: analysis; carbon sources; gluconic acid: secondary carbon source

ORGN .

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Microorganisms; Irregular Nonsporing Gram-Positive Rods: Actinomycetes
        and Related Organisms, Eubacteria, Bacteria, Microorganisms;
        Microorganisms
ORGN Organism Name
        Bacillus subtilis (Endospore-forming Gram-Positives);
        Corynebacterium glutamicum (Irregular Nonsporing Gram-Positive
        Rods): non-growing cells; Escherichia coli (Enterobacteriaceae);
        bacteria (Bacteria); microorganisms (Microorganisms)
ORGN Organism Superterms
        Bacteria; Eubacteria; Microorganisms
     9001-82-5Q (6-PHOSPHOGLUCONATE DEHYDROGENASE)
RN
     9073-95-4Q (6-PHOSPHOGLUCONATE DEHYDROGENASE)
     56-87-1 (L-LYSINE)
     526-95-4 (GLUCONIC ACID)
L15 ANSWER 2 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS
                    2001:174925 BIOSIS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                    PREV200100174925
TITLE:
                    MALDI-TOF MS for quantification of substrates and products
                    in cultivations of Corynebacterium glutamicum.
AUTHOR(S):
                    Wittmann, Christoph (1); Heinzle, Elmar
CORPORATE SOURCE:
                    (1) Biochemical Engineering Institute, Saarland
University,
                    66041, Saarbruecken: c.wittmann@rz.uni-sb.de Germany
SOURCE:
                    Biotechnology and Bioengineering, (March 20, 2001) Vol.
72,
                    No. 6, pp. 642-647. print.
                    ISSN: 0006-3592.
DOCUMENT TYPE:
                    Article
LANGUAGE:
                    English
SUMMARY LANGUAGE:
                    English
    MALDI-TOF MS for quantification of substrates and products in
cultivations
     of Corynebacterium glutamicum.
     The application of MALDI-TOF MS for the quantification of lysine
AB
     , alanine, and glucose is described. The method is based on using stable
     isotopes as internal standards and allows fast, sensitive,. .
     concentrations of the analytes between 10 muM and 100 mM. The mean
     standard deviations from five replicates each were 4.3% (lysine
     ), 3.7% (alanine), and 3.2% (glucose). In addition, sucrose could be
     measured by MALDI-TOF MS, but was not quantified due to.
TΤ
     Major Concepts
        Biochemistry and Molecular Biophysics; Bioprocess Engineering; Methods
        and Techniques
TT
     Chemicals & Biochemicals
          amino acids: microbial production
        , quantitative analysis; products: quantitative analysis; substrates:
        quantitative analysis
L15 ANSWER 3 OF 13 CAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER:
                         2000:761605 CAPLUS
DOCUMENT NUMBER:
                         134:99608
TITLE:
                         Development and use of miniaturized parallel
                         experiment technology for bioprocess development
AUTHOR(S):
                         Altenbach-Rehm, Jutta
```

Institut fur Biotechnologie, Julich, JUL-3782,

Ber. Forschungszent. Juelich (2000), Juel-3782, a-f,

CORPORATE SOURCE:

Germany SOURCE:

i-iv, 1-233

CODEN: FJBEE5; ISSN: 0366-0885

DOCUMENT TYPE: LANGUAGE:

Report German

AB The fed-batch technique is nowadays the std. operation mode for high performance microbial prodn. processes. Shake flasks are widely used a simple bioreactors in batch process development. Because of uncontrolled changes in pH and reduced oxygen transfer rates parallel operated shake flasks can usually not be applied for microbial fed-batch process development. Due to the need to operate controlled stirred tank reactors the development of specific fermn. strategies is expensive and time consuming. To address these issues a miniature fed-batch technique was developed in cooperation with DASGIP mbH, Julich, and INFORS AG, Basel. Controlled batch and fed-batch fermns. can be performed in 16 parallel small scale bioreactors. The new technique allows the feeding of up to 4 different substrates and parallel pH control

based on user-defined profiles. The feeding assembly is sterilized chem. using dimethylcarbonate. To overcome the limitations of shake flasks, small scale bubble columns were developed. Gas distribution is performed with a new type of sterile sparger. Transferring fed-batch fermns. from small scale bubble columns to a stirred tank reactor a scale up factor of 20 was achieved. Escherichia coli K12 was chosen to test the new parallel

bioreactor technique. Compared to shake flask fermns, the cell concn. was

50% higher due to efficient oxygen transfer. In fed-batch fermns. with pH-controlled substrate feeding up to 35 g/l DCW were achieved. For the 1st time, a parallel optimization of feeding profiles in parallel small scale fed-batch expts. with successful scale-up to a lab. bioreactor was performed. Escherichia coli BL 21 (DE3) pLysS produces the recombinant enzyme GDP-.alpha.-D-mannose-pyrophosphorylase after induction with isopropyl-.beta.-D-thiogalactoside (IPTG). Single induction with 0,5 mM IPTG resulted in a low specific enzyme activity of 1,6 U/g DCW (dry cell wt.). For the optimization of enzyme expression a genetic algorithm was used. A final enzyme activity of 111 U/g DCW was achieved for optimal substrate and inducer feeding profiles. To demonstrate the advantages of this new parallel bioreactor technique different strains of industrial relevance were investigated. Corynebacterium glutamicum, Staphylococcus carnosus and Ashbya gossypii.

ST bioreactor bubble column fed batch fermn; GDP mannose pyrophosphorylase bubble column fed batch; Staphylococcus bubble column fed batch fermn; isoleucine bubble column fed batch Corynebacterium; riboflavin bubble column fed batch Ashbya

IT Amino acids, biological studies

RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (amino acid consumption in riboflavin prodn. by Ashybya gossypii in parallel bubble columns with fed-batch technique)

IT Corynebacterium glutamicum

(L-isoleucine prodn. by **Corynebacterium** glutamicum in parallel bubble columns with fed-batch technique)

IT 56-40-6, Glycine, biological studies 56-41-7, Alanine, biological studies 56-45-1, Serine, biological studies 56-84-8, Aspartic acid, biological studies 56-85-9, Glutamine, biological studies 56-86-0, Glutamic acid, biological studies 56-87-1, Lysine, biological studies 60-18-4, Tyrosine, biological studies 63-68-3, Methionine, biological studies 63-91-2, Phenylalanine, biological studies 72-18-4.

Valine, biological studies 72-19-5, Threonine, biological studies 73-22-3, Tryptophane, biological studies RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (amino acid consumption in riboflavin prodn. by Ashybya gossypii in parallel bubble columns with fed-batch technique) 73-32-5P, L-Isoleucine, biological studies TΤ RL: BMF (Bioindustrial manufacture); BPR (Biological process); BIOL (Biological study); PREP (Preparation); PROC (Process) (L-isoleucine prodn. by Corynebacterium glutamicum in parallel bubble columns with fed-batch technique, amino acid consumption in riboflavin prodn. by Ashybya gossypii) IT 61-90-5, L-Leucine, biological studies RL: BPR (Biological process); MFM (Metabolic formation); BIOL (Biological study); FORM (Formation, nonpreparative); PROC (Process) (L-isoleucine prodn. by Corynebacterium glutamicum in parallel bubble columns with fed-batch technique, amino acid consumption in riboflavin prodn. by Ashybya gossypii) L15 ANSWER 4 OF 13 CAPLUS COPYRIGHT 2001 ACS ACCESSION NUMBER: 1999:244776 CAPLUS DOCUMENT NUMBER: 130:266420 TITLE: Method for microbial production of amino acids of the aspartate and/or glutamate family and agents which can be used in said method INVENTOR(S): Eikmanns, Bernd; Peters-Wendisch, Petra; Sahm, Hermann PATENT ASSIGNEE(S): Forschungszentrum Julich G.m.b.H., Germany PCT Int. Appl., 40 pp. SOURCE: CODEN: PIXXD2 DOCUMENT TYPE: Patent LANGUAGE: German FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE --------------WO 9918228 A2 19990415 WO 1998-EP6210 19980930 WO 9918228 A3 19990520 W: AU, BR, CA, CN, HU, ID, JP, KR, MX, RU, SK, US RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE DE 19831609 Α1 19990415 DE 1998-19831609 19980714 AU 9911482 AU 1999-11482 A1 19990427 19980930 EP 1015621 EP 1998-954301 A2 20000705 19980930 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI BR 9813021 Α 20000815 BR 1998-13021 19980930 PRIORITY APPLN. INFO.: DE 1997-19743894 A 19971004 DE 1998-19831609 A 19980714 WO 1998-EP6210 W 19980930 TI Method for microbial production of amino acids of the aspartate and/or glutamate family and agents which can be used in said method AB The invention relates to a method for microbial prodn. of amino acids of the aspartate and/or glutamate family in which the pyruvate carboxylase activity is increased by genetically changing the enzyme and/or the pyruvate carboxylase gene

expression of a microorganism which produces the corresponding amino acid. In addn., the invention relates to a pyruvate carboxylase gene and addnl. agents which can be used in the inventive method. ST amino acid fermn Corynebacterium pyruvate carboxylase genetic engineering ΙT Corynebacterium glutamicum Fermentation (microbial prodn. of amino acids of the aspartate and/or glutamate family and agents which can be used in said method) IT Amino acids, preparation RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (microbial prodn. of amino acids of the aspartate and/or glutamate family and agents which can be used in said method) IT Genetic engineering (microbial prodn. of amino acids of the aspartate and/or glutamate family and modification of Corynebacterium pyc gene in said method) ΙT Genes (microbial) RL: BPR (Biological process); BIOL (Biological study); PROC (Process) (pyc; microbial prodn. of amino acids of the aspartate and/or glutamate family and modification of Corynebacterium pyc gene in said method) TΨ 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, 72-19-5P, L-Threonine, preparation preparation 74-79-3P, L-Arginine, 672-15-1P, L-Homoserine preparation RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (microbial prodn. of amino acids of the aspartate and/or glutamate family and agents which can be used in said method) ΤТ 9014-19-1, Pyruvate carboxylase 204116-18-7 208541-81-5 RL: BSU (Biological study, unclassified); BIOL (Biological study) (microbial prodn. of amino acids of the aspartate and/or glutamate family and agents which can be used in said method) L15 ANSWER 5 OF 13 BIOSIS COPYRIGHT 2001 BIOSIS ACCESSION NUMBER: 1999:175414 BIOSIS DOCUMENT NUMBER: PREV199900175414 TITLE: Cloning of the transketolase gene and the effect of its dosage on aromatic amino acid production in Corynebacterium glutamicum.

AUTHOR(S): Ikeda, M. (1); Okamoto, K.; Katsumata, R.

CORPORATE SOURCE: (1) Technical Research Laboratories, Kyowa Hakko Kogyo

Co.,

Ltd., Hofu, Yamaguchi, 747-8522 Japan

Applied Microbiology and Biotechnology, (Feb., 1999) Vol. SOURCE:

51, No. 2, pp. 201-206.

ISSN: 0175-7598.

DOCUMENT TYPE: Article LANGUAGE: English

TТ Cloning of the transketolase gene and the effect of its dosage on aromatic

amino acid production in Corynebacterium glutamicum.

 ${\tt AB.}$. . enzyme of the non-oxidative pentose phosphate pathway. The effect of

its overexpression on aromatic amino acid production was investigated in <code>Corynebacterium</code> glutamicum, a typical amino-acid-producing organism. For this purpose, the transketolase gene of the organism was cloned on the basis of. . . as a protein of approximately 83kDa in proportion to the copy numbers. Introduction of the plasmids into a tryptophan and <code>lysine</code> co-producer resulted in copy-dependent increases in tryptophan production along with concomitant decreases in <code>lysine</code> production. Furthermore, the presence of the gene in high copy numbers enabled tyrosine, phenylalanine and tryptophan producers to accumulate 5%-20% more aromatic <code>amino acids</code>. These results indicate that overexpressed transketolase activity operates to redirect the glycolytic intermediates toward the nonoxidative pentose phosphate pathway in. . .

IT Major Concepts

Bioprocess Engineering; Enzymology (Biochemistry and Molecular Biophysics); Molecular Genetics (Biochemistry and Molecular Biophysics)

IT Chemicals & Biochemicals

aromatic amino acids: microbial

production; transketolase [EC 2.2.1.1]; Corynebacterium
transketolase gene (Irregular Nonsporing Gram-Positive Rods)

L15 ANSWER 6 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1995:674124 CAPLUS

DOCUMENT NUMBER:

123:54314

TITLE:

Enhancement of reduced NADP production for enhanced

microbial production of biochemicals Kojima, Hiroyuki; Totsuka, Kazuhiko

PATENT ASSIGNEE(S):

Ajinomoto Co., Inc., Japan

SOURCE:

PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

INVENTOR(S):

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

r: 1

PATENT INFORMATION:

	PATENT NO.			KIND DATE				APPLICATION NO.							DATE				
	WO	9511	985		A	1	 1995	0504		1	wo 1	994-	JP17	791	-	1994	1026		
		W:	AU,	BR,	CA,	CN,	CZ,	HU,	JP,	KR,	, PI	, RU	, SI	ζ, ί	JS,	VN			
		RW:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	, GF	R, IE	, II	., I	ŪŪ,	MC,	NL,	PT,	SE
	CA	2175	042		A	Ą	1995	0504		(CA 1	994-	2175	042	2	1994	1026		
	AU	9480	026		A	1	1995	0522		7	AU 1	994-	8002	26		1994	1026		
	EΡ	7337	12		A	1	1996	0925]	EP 1	994-	9311	.58		1994	1026		
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	, GF	R, IE	, I	, I	Ί,	LU,	MC,	NL,	PT,
SE																			
	BR	9407	907		Α		1996	1126		1	3R 1	994-	7907	7		1994	1026		
	HU	7484	0		A:	2	1997	0228		I	HU 1	.996-	1085	5		1994	1026		
	zA	9503	350		Α		1996	1025		2	ZA 1	.995-	3350)		1995	0425		
	US	5830	716		Α		1998	1103		Ţ	JS 1	.996-	6195	21		1996	0429		
	CN	1139	956		Α		1997	0108		(CN 1	994-	1947	707		1996	1026		
PRIO	RITY	APP	LN.	INFO	. :					JP :	1993	3-270	828			1993	1028		
									1	WO :	1994	-JP1	791			1994	1026		

TI Enhancement of reduced NADP production for enhanced microbial production of biochemicals

The productivity of such substances as L-amino acids, antibiotics, vitamins, growth factors and physiol. active substances in the fermn. using a microorganism is improved by improving the productivity of reduced NADP in the cells of the microorganisms. Construction of pMW::THY contg. the Escherichia coli transhydrogenase gene, and introduction of the plasmid into the L-threonine-producing Escherichia coli B-3996 were shown. The recombinant E. coli B-3996 produced L-threonine .apprx.10% higher than did the parental strain. TΤ Corynebacterium glutamicum Escherichia coli Fermentation (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) TΤ Amino acids, preparation RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) ΙT Plasmid and Episome (pHSG::THY; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) Plasmid and Episome ΙT (pMW::THY; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) IT Plasmid and Episome (pSU::THY; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) TT 9014-18-0, Nicotinamide nucleotide transhydrogenase RL: BAC (Biological activity or effector, except adverse); BIOL (Biological study) (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) 56-86-0P, L-Glutamic acid, preparation 56-87-1P, L-Lysine, preparation 61-90-5P, L-Leucine, preparation 63-91-2P, L-Phenylalanine, preparation 72-18-4P, L-Valine, preparation 72-19-5P, L-Threonine, preparation 73-32-5P, L-Isoleucine, preparation RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) IT 53-59-8P, NADP RL: BPN (Biosynthetic preparation); BIOL (Biological study); PREP (Preparation) (reduced; enhancement of reduced NADP prodn. for enhanced microbial prodn. of biochems.) L15 ANSWER 7 OF 13 SCISEARCH COPYRIGHT 2001 ISI (R) ACCESSION NUMBER: 95:184306 SCISEARCH THE GENUINE ARTICLE: QK574 METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM TITLE: CORYNEBACTERIUM-GLUTAMICUM AUTHOR: SAHM H (Reprint); EGGELING L; EIKMANNS B; KRAMER R CORPORATE SOURCE: KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL, D-52425 JULICH, GERMANY (Reprint) COUNTRY OF AUTHOR:

FEMS MICROBIOLOGY REVIEWS, (FEB 1995) Vol. 16, No. 2-3,

GERMANY

SOURCE:

pp. 243-252.

ISSN: 0168-6445.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT:

ENGLISH

LANGUAGE:

LIFE

REFERENCE COUNT:

36

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS METABOLIC DESIGN IN AMINO-ACID PRODUCING BACTERIUM CORYNEBACTERIUM

-GLUTAMICUM

AB The Gram-positive bacterium Corynebacterium glutamicum is used for the industrial production of amino acids, e.g. of L-glutamate and L-lysine. In the last 10 years, genetic engineering and amplification of relevant structural genes have become fascinating methods for the construction of strains with desired genotypes. By cloning and expressing the various genes of the Llysine pathway in C. glutamicum we could demonstrate that an increase of the flux of L-aspartate semialdehyde to L-lysine could be obtained in strains with increased dehydrodipicolinate synthase activity. By combined overexpression of deregulated aspartate kinase and dihydrodipicolinate synthase, the L-lysine secretion could be increased (10-20%). Recently we detected that in C. glutamicum two pathways exist for the synthesis of DL-diaminopimelate and Llysine. Mutants defective in one pathway are still able to synthesize enough L-lysine for growth, but the L-lysine secretion is reduced to 50-70%. Using NMR spectroscopy, we could

calculate

how much of the L-lysine secreted into the medium is synthesized via each pathway. Amplification of the feedback inhibition-insensitive homoserine dehydrogenase and homoserine kinase in a high L-lysine overproducing strain enabled channelling of the carbon flow from the intermediate aspartate semialdehyde towards homoserine, resulting in a high accumulation. . . acid overproduction, the secretion into the culture medium also has to be noted. Recently it could be demonstrated that L-glutamate, L-lysine and L-isoleucine are not secreted via passive diffusion but via specific active carrier systems. Analysis of lysine-overproducing C. glutamicum strains indicates that this secretion carrier has a strong influence on the overproduction of this amino acid. Thus,.

ST Author Keywords: CORYNERACTERIUM GLUTAMICUM; AMINO ACID PRODUCTION; METABOLIC DESIGN; L-LYSINE; L-THREONINE; L-ISOLEUCINE

KeyWords Plus (R): L-THREONINE; L-LYSINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; MICROBIAL PRODUCTION; RESISTANT MUTANTS; SPLIT PATHWAY; BIOSYNTHESIS; ISOLEUCINE; GENES

L15 ANSWER 8 OF 13 SCISEARCH COPYRIGHT 2001 ISI (R)

ACCESSION NUMBER: 96:7187 SCISEARCH

THE GENUINE ARTICLE: TJ545

TITLE:

METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM

CORYNEBACTERIUM-GLUTAMICUM

AUTHOR:

SAHM H (Reprint)

CORPORATE SOURCE:

KFA JULICH GMBH, FORSCHUNGSZENTRUM, INST BIOTECHNOL,

D-52425 JULICH, GERMANY (Reprint)

COUNTRY OF AUTHOR:

GERMANY

SOURCE:

FOLIA MICROBIOLOGICA, (1995) Vol. 40, No. 1, pp. 23-30.

ISSN: 0015-5632.

DOCUMENT TYPE:

Article; Journal

FILE SEGMENT:

LIFE; AGRI

LANGUAGE:

ENGLISH

REFERENCE COUNT:

26

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

TI METABOLIC DESIGN IN THE AMINO-ACID-PRODUCING BACTERIUM

CORYNEBACTERIUM-GLUTAMICUM

The Gram-positive bacterium Corynebacterium glutamicum is used for the industrial production of amino acids, e.g. of L-glutamate and L-lysine. By cloning and expressing the various genes of the L-lysine pathway in C. glutamicum we could demonstrate that an increase of the flux of L-4-aspartaldehydate to L-lysine could be obtained in strains with increased dihydro-dipicolinate synthase activity. Recently we detected that in C. glutamicum two pathways exist for the synthesis of DL-2,6-diaminopimelate and L-lysine. Mutants defective in one pathway are still able to synthesize enough L-lysine for growth but the L-lysine

secretion is reduced to 50-70 %. Using NMR-spectroscopy we could calculate

how much of the L-lysine secreted into the medium is synthesized via the one and the other pathway. Amplification of the feedback-inhibition-insensitive-homoserine dehydrogenase and homoserine kinase in a high L-lysine-overproducing strain made it possible to channell of the carbon now from the intermediate 4-aspartaldehydate toward homoserine, resulting in a high. . .

STP KeyWords Plus (R): L-THREONINE; BREVIBACTERIUM-LACTOFERMENTUM; HOMOSERINE DEHYDROGENASE; MICROBIAL PRODUCTION; LYSINE BIOSYNTHESIS; RESISTANT MUTANTS; SPLIT PATHWAY; GENES; AMPLIFICATION; FERMENTATION

L15 ANSWER 9 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1985:436156 CAPLUS

DOCUMENT NUMBER:

103:36156

TITLE:

Optimization of amino acid production by automatic

self-tuning digital control of redox potential Radjai, Mohammad K.; Hatch, Randolph T.; Cadman,

Theodore W.

CORPORATE SOURCE:

Dep. Chem. Nucl. Eng., Univ. Maryland, College Park,

MD, 20742, USA

SOURCE:

Biotechnol. Bioeng. Symp. (1984), 14 (Symp.

Biotechnol.

AUTHOR(S):

Fuels Chem., 6th), 657-79

CODEN: BIBSBR; ISSN: 0572-6565

DOCUMENT TYPE:

Journal English

LANGUAGE:

The microbial prodn. of homoserine [672-15-1],

lysine [56-87-1], and valine [72-18-4] by an auxotrophic mutant of Corynebacterium glutamicum was investigated in a 16-L batch fermentor. Closed-loop digital control of redox potential was implemented

using proportional-integral (PI) control of agitation rates. Due to the nonlinearity of the system, the PI controller parameters had to be changed

during the course of the fermns. An automatic, self-tuning algorithm was developed for stable control of redox potential. This permitted exptl. optimization of total and selective amino acid prodn. Total amino acid yields of 35% from glucose were achieved compared to 23% reported in the literature for the same fermn.

ST amino acid fermn redox potential control; optimization simulation

```
homoserine lysine valine fermn
IT
     Corynebacterium glutamicum
        (amino acid manuf. with, optimization and redox potential control in)
ΤТ
     Amino acids, preparation
     RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP
     (Preparation)
        (manuf. of, by fermn.)
L15 ANSWER 10 OF 13
                         MEDLINE
ACCESSION NUMBER:
                    79079819
                                 MEDLINE
DOCUMENT NUMBER:
                    79079819
                               PubMed ID: 727028
TITLE:
                    Microbial production of essential amino
                    acid with Corynebacterium glutamicum mutants.
AUTHOR:
                    Nakayama K; Araki K; Kase H
SOURCE:
                    ADVANCES IN EXPERIMENTAL MEDICINE AND BIOLOGY, (1978) 105
                    649-61.
                    Journal code: 2LU; 0121103. ISSN: 0065-2598.
PUB. COUNTRY:
                    United States
                    Journal; Article; (JOURNAL ARTICLE)
LANGUAGE:
                    English
FILE SEGMENT:
                    Priority Journals
ENTRY MONTH:
                    197902
ENTRY DATE:
                    Entered STN: 19900314
                    Last Updated on STN: 19970203
                    Entered Medline: 19790212
ΤI
     Microbial production of essential amino acid with
     Corynebacterium glutamicum mutants.
     Amino acids produced by microbial process are
AB
     generally L-forms. The stereospecificity of the amino
     acids produced by fermentation makes the process advantageous
     compared with synthetic process. Microorganisms employed in microbial
     process for amino acid production are divided into 4 classes; wild-type
     strain, auxotrophic mutant, regulatory mutant and auxotrophic regulatory
     mutant. Using such mutants of Corynebacterium glutamicum, all
     the essential amino acids but L-methionine are now
     being produced by "direct fermentation" from cheap carbon sources such as
     carbohydrate materials or acetic acid.
СТ
     *Amino Acids, Essential: BI, biosynthesis
       *Corynebacterium: ME, metabolism
      Fermentation
      Kinetics
      Leucine: BI, biosynthesis
        Lysine: BI, biosynthesis
      Mutation
      Phenylalanine: BI, biosynthesis
      Species Specificity
      Stereoisomerism
      Threonine: BI, biosynthesis
      Tryptophan: BI, biosynthesis
     3617-44-5 (Phenylalanine); 56-87-1 (Lysine); 7005-03-0
RN
     (Leucine); 72-19-5 (Threonine); 73-22-3 (Tryptophan)
CN
     0 (Amino Acids, Essential)
L15 ANSWER 11 OF 13 CAPLUS COPYRIGHT 2001 ACS
ACCESSION NUMBER:
                         1970:508239 CAPLUS
DOCUMENT NUMBER:
                         73:108239
TITLE:
```

Microbial production of

L-threonine

INVENTOR(S):

Nakayama, Kiyoshi; Kase, Hiroshi

PATENT ASSIGNEE(S):

Kyowa Fermentation Industry Co. Ltd.

SOURCE:

Ger. Offen., 22 pp. CODEN: GWXXBX

DOCUMENT TYPE:

Patent

LANGUAGE:

FAMILY ACC. NUM. COUNT:

German

PATENT INFORMATION:

PATENT NO.

KIND DATE

APPLICATION NO. DATE

DE 1817666

Α 19700827 DE 1968-1817666 19681224

ΤТ Microbial production of L-threonine

AB Various microorganisms, e.g. Aerobacter [Enterobacter] aerogenes, Serratia

marcescens, or Arthrobacter paraffineus, cultured for producing L-threonine required 2 or 3 of the amino acids isoleucine, methionine, lysine, or diaminopimelic acid. The microorganisms were cultured aerobically in an aq. medium contg. the optimal (or less) amts. of the required amino acids. Thus, E. aerogenes NM-IS-5 (ATCC 21,215) was cultured 96 hr at 30.degree. in medium contg. glucose 5, (NH4)2SO4 1.4, KH2PO4 0.05, K2HPO4 0.05, MgSO4.7H2O 0.025, FeSO4.7H2O 0.001, MnSO4.4H2O 0.001, and CaCO3 2% and isoleucine 50, methionine 100, and diaminopimelic acid 200 mg/l. to give 7.8 g L-threonine/l.

ST microbial prodn threonine; threonine microbial

prodn; Aerobacter threonine fermn; amino acid prodn fermn

TT Corynebacterium

(glutamicum, threonine manuf. by)

L15 ANSWER 12 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1967:514494 CAPLUS

DOCUMENT NUMBER:

67:114494

TITLE:

Microbial production of

amino acids from hydrocarbons. III.

L-Ornithine production by an arginine auxotrophic

mutant of Corynebacterium hydrocarboclastus Ishu, Ryosuke; Ishii, Ryosuke; Shiio, Isamu

AUTHOR(S):

Ajinomoto Co., Inc., Kawasaki, Japan

CORPORATE SOURCE: SOURCE:

J. Gen. Appl. Microbiol. (1967), 13(3), 3303-12

CODEN: JGAMA9

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Microbial production of amino acids

from hydrocarbons. III. L-Ornithine production by an arginine auxotrophic mutant of Corynebacterium hydrocarboclastus

AB cf. CA 67: 89718u. The arginine auxotrophic mutant strain RN-362 of C. hydrocarboclastus R-7 was used to study L-ornithine productiion from hydrocarbons, in a fermentation medium contg. various n-alkanes. L-Ornithine production required L-arginine at the optimum level of

g./l. of medium; an excess inhibited the biosynthesis of L-ornithine. (NH4)2HPO4 was the best source of N and, at 2% in a neutral to slightly acidic pH, gave the highest level of L-ornithine production and cell growth; NH4OAc, KNO3, and (NH4)2CO3 proved less suitable because of a

drop

in pH along with the accumulation of a large amt. of .alpha.-ketoglutaric acid, pyruvic acid, and proline in the growth medium. Of 17 C sources,

n-tetradecane best supported cell growth and L-ornithine production and the other C13-C17 n-alkanes did so moderately, while kerosene and light oil produced good cell growth but only a small amt. of L-ornithine.

Addn.

of 3 g. yeast ext. and 0.5 g. L-arginine-HCl to 1 l. of medium enhanced L-ornithine production. A similar effect was achieved by replacing the yeast ext. with various amino acids at 0.01% in the medium. L-Methionine was most effective for the production of L-ornithine, while L-lysine, L-cysteine, L-cystine, L-histidine, and L-phenylalanine were less so, in decreasing order. Amino acids enhance L-ornithine production by stimulating hydrocarbon oxidn. and cell growth.

ST HYDROCARBONS USE BACTERIA; BACTERIA HYDROCARBONS USE; ALKANES USE BACTERIA; AMINO ACIDS PRODN HYDROCARBONS; ORNITHINE PRODN HYDROCARBONS; PARAFFINS UTILIZATION BACTERIA

TT Corvnebacterium

(hydrocarboclastus, ornithine formation from hydrocarbons by)

Hydrocarbons, biological studies IT

RL: BIOL (Biological study)

(ornithine formation from, by Corynebacterium

hydrocarboclastus)

IT 70-26-8

RL: FORM (Formation, nonpreparative)

(formation of, from hydrocarbons by Corynebacterium hydrocarboclastus)

L15 ANSWER 13 OF 13 CAPLUS COPYRIGHT 2001 ACS

ACCESSION NUMBER:

1967:489718 CAPLUS

DOCUMENT NUMBER:

67:89718

TITLE:

Microbial production of

amino acids from hydrocarbons. II.

Isolationf good hycarbon utilizers and amino acid

production by their auxotrophs

AUTHOR(S):

Ishii, Ryosuke; Otsuka, Shinichiro; Shiio, Isamu CORPORATE SOURCE: Central Res. Labs., Ajinomoto Co., Inc., Kawasaki,

SOURCE:

J. Gen. Appl. Microbiol. (1967), 13(2), 217-25

CODEN: JGAMA9

DOCUMENT TYPE:

Journal

LANGUAGE:

English

Microbial production of amino acids

from hydrocarbons. II. Isolationf good hycarbon utilizers and amino acid

production by their auxotrophs

- AΒ cf. CA 59: 14313h. Nine microorganisms, which showed good growth on long-chain aliphatic hydrocarbons, were isolated by an enrichment culture method, followed by a single colony isolation technique. They included 5 strains of Alcaligenes marshallii, 2 strains of Corynebacterium hydrocarboclastus, and 2 strains of yeast. Various auxotrophic mutants were derived from these microorganisms. The mutants accumulated the following amino acids from aliphatic hydrocarbons; L-ornithine, L-valine, L-glutamic acid, L-leucine, L-tyrosine, L-alanine, L-proline, L-aspartic acid, and L-lysine.
- STBACTERIA AMINO ACID PRODN; AMINO ACID PRODN BACTERIA; HYDROCARBONS AMINO ACIDS; ALIPHATICS BACTERIA METAB
- ITCorynebacterium

(hydrocarboclastus, amino acid fermentation of hydrocarbons by)

IT Amino acids, preparation

```
(Preparation)
        (manuf. of, by fermentation of hydrocarbons)
=> dis his
     (FILE 'HOME' ENTERED AT 18:56:31 ON 19 OCT 2001)
     FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE, SCISEARCH' ENTERED AT 18:57:06 ON
     19 OCT 2001
L1
           2893 S MICROBIAL (W) PRODUCTION
L2
            190 S L1 AND (AMINO (W) ACIDS)
L3
             28 S L2 AND CORYNEBACTERIUM
L4
             13 L3 AND LYSINE
L5
              0 L3 AND ((EXPORT) (W) (GENE OR CARRIER))
L6
              0 S L3 AND EXPORT (W) GENE
            166 S EXPORT (W) GENE
L7
L8
              0 S L3 AND L7
L9
              0 S L3 (P) L7
L10
              0 S L2 AND L7
              0 S L2 AND EXPORT (W) GENE
L11
L12
             61 S L7 AND MICROB?
L13
             1 S L12 AND CORYNEBACTERIUM
L14
             26 DUP REM L3 (2 DUPLICATES REMOVED)
L15
             13 DUP REM L4 (0 DUPLICATES REMOVED)
=> log off y
```

RL: BMF (Bioindustrial manufacture); BIOL (Biological study); PREP

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STN INTERNATIONAL LOGOFF AT 19:32:29 ON 19 OCT 2001